



Cancer Research

**American Association for
Cancer Research, Inc.**

Publications Staff

Managing Editor
Margaret Foti

Assistant Managing Editor
Mary Anne Mennite

Production Manager
Heide M. Pusztay

Staff Editor
Martha Michael

Editorial Assistants
Angela Barry
Linda Pote Musumeci
James D. Tang

**Supervisor, Manuscript
Processing**
Margaret A. Pickels

**Assistant Supervisor,
Manuscript Processing**
Theresa A. Griffith

Editorial Secretaries
Mary Ellen Piring
June A. Holloway
Valerie L. Taylor
Luz M. Torres

Administrative Staff
Executive Director
Margaret Foti

**Associate Director of
Administration**
Adam D. Blistein

Senior Accountant
Anthony P. Blengino

Financial Assistant
George L. Moore

**Meetings and Development
Coordinator**
Jeffrey M. Ruben

**Executive Assistant for
Programs**
Ruth E. Fortson

Administrative Assistant
Elizabeth A. Moore

**Staff Assistant for Member
Services**
Robin E. Felder

Editorial Secretary
Diana F. Certo

Secretary
Shirley L. Huestis

Editor-in-Chief
Carlo M. Croce

Associate Editors

Stuart A. Aaronson

Jerry M. Adams

David S. Alberts

William M. Baird

Charles M. Balch

Allan Balmain

J. Carl Barrett

Renato Baserga

William T. Beck

Thomas L. Benjamin

June L. Biedler

Peter M. Blumberg

Gianni Bonadonna

G. Tim Bowden

Luther W. Brady

Norman Breslow

Edward Bresnick

Samuel Broder

Sam C. Brooks

Ronald N. Buick

C. Patrick Burns

Bruno Calabretta

Eli Canaani

Robert L. Capizzi

Bruce A. Chabner

Yung-chi Cheng

David Colcher

Allan H. Conney

Neal G. Copeland

Joseph G. Cory

Jeffrey Cossman

Frank Costantini

Benoit de Crombrughe

Riccardo Dalla-Favera

Richard L. Davidson

Eugene R. DeSombre

T. Michael Dexter

Ralph E. Durand

Alan Eastman

Gertrude B. Elion

Lois B. Epstein

Leonard C. Erickson

M. Essex

Emmanuel Farber

Nelson Fausto

Isaiah J. Fidler

Robert A. Floyd

Judah Folkman

Kenneth A. Foon

Joseph F. Fraumeni, Jr.

Errol C. Friedberg

Minoru Fukuda

Philip Furmanski

Eugene W. Gerner

Leo E. Gerweck

Eli Glatstein

David W. Golde

Michael M. Gottesman

J. W. Grisham

Sen-itiroh Hakomori

Philip C. Hanawalt

Curtis C. Harris

Jules E. Harris

John C. Harshbarger

Stephen S. Hecht

Ingegerd Hellstrom

Brian E. Henderson

Gloria H. Heppner

Russell Hilf

Richard B. Hochberg

Tasuku Honjo

Kay Huebner

James N. Ihle

John T. Isaacs

Mark A. Israel

Rakesh K. Jain

Peter A. Jones

V. Craig Jordan

Robert S. Kerbel

Young S. Kim

Tadamitsu Kishimoto

Alfred G. Knudson, Jr.

Kurt W. Kohn

Laurence N. Kolonel

Donald W. Kufe

John S. Lazo

Brigid G. Leventhal

Jay A. Levy

Frederick P. Li

Michael W. Lieberman

Victor Ling

Lance A. Liotta

Martin Lipkin

Marc E. Lippman

Gerald Litwack

Leroy F. Liu

Lawrence A. Loebe

Dan L. Longo

Diana M. Lopez

Reuben Lotan

James W. Lown

David B. Ludlum

Tak W. Mak

Michael J. Mastrangelo

W. Gillies McKenna

Anna T. Meadows

Myron R. Melamed

John Mendelsohn

Christopher J. Michejda

John D. Minna

Malcolm S. Mitchell

Harold L. Moses

Sharon B. Murphy

Garth L. Nicolson

Kenneth Nilsson

Jeffrey A. Norton

Peter C. Nowell

Kenneth Olden

Gilbert S. Omenn

Richard J. O'Reilly

Robert F. Ozols

Michael A. Palladino, Jr.

Peter L. Pedersen

Anthony E. Pegg

Angel Pellicer

Bice Perussia

Sidney Pestka

Cecil B. Pickett

W. J. Pledger

Frank J. Rauscher III

Donald J. Reed

Ralph A. Reisfeld

Richard A. Rifkind

Robert T. Schimke

J. Schlessinger

Manfred Schwab

Joseph V. Simone

Beatrice Singer

Francis M. Sirotnak

Arthur A. Spector

Martha R. Stampfer

Eric J. Stanbridge

Gary S. Stein

Bernard S. Strauss

Takashi Sugimura

Saraswati Sukumar

James A. Swenberg

Paul Talalay

James E. Talmadge

Masaaki Terada

Snorri S. Thorgeirsson

Donald J. Tindall

George J. Todaro

Jeffrey M. Trent

Giorgio Trinchieri

Axel Ullrich

Peter Vaupel

Ellen S. Vitetta

Daniel D. Von Hoff

Peter A. Ward

Michael D. Waterfield

Lee W. Wattenberg

I. Bernard Weinstein

Bengt Westermark

Raymond L. White

Gordon F. Whitmore

Walter Willett

Sheldon Wolff

Charles W. Young

Stuart H. Yuspa

Harald zur Hausen

Cover Editorial Board

Sidney Weinhouse,
Cover Editor

Joseph H. Burchenal
Nancy H. Colburn

Hugh J. Creech
George H. Hitchings

Edwin A. Mirand
John H. Weisburger

Takashi Sugimura

Cancer Research is sponsored by the American Association for Cancer Research, Inc. and receives major support from the American Cancer Society, Inc. Publication costs are also met by a grant from the Elsa U. Pardee Foundation.

Subscription Information

Cancer Research is published twice a month, one volume per year, by the American Association for Cancer Research, Inc. Subscriptions include the *Proceedings of the American Association for Cancer Research*, issued in April of each year. Except for members of the American Association for Cancer Research, all subscriptions are payable in advance to Waverly Press, Inc., 428 E. Preston Street, Baltimore, MD 21202-3993, to which all business communications, remittances (in United States currency or its equivalent), and subscription orders should be sent. In Japan, send orders and inquiries to (sole agent): USACO Corporation, Tsutsumi Bldg., 13-12, Shimbashi-chome, Minato-ku, Tokyo 105, Japan; Tel. (03) 502-6471. Nonmembers of the American Association for Cancer Research within the United States may subscribe to Volume 52 (1992) of *Cancer Research* at the rate of \$350.00 (individual and institutional nonmembers), including postage. Institutional and individual nonmembers outside of the United States should add \$60.00 to their respective rates to offset second-class postage costs. Air mail and other means of delivery are available at extra charge; contact Waverly Press for information. Changes of address notification should be sent 60 days in advance and include both old and new addresses. Member subscribers should send address changes to: Member Services, American Association for Cancer Research, Inc., Public Ledger Building, 620 Chestnut St., Suite 816, Philadelphia, PA 19106-3483. Nonmember subscribers should notify Waverly Press of changes of address. Copies of the journal which are undeliverable because of address changes will be destroyed.

Notice to Members of the American Association for Cancer Research

Officers for 1991-1992

President: Harold L. Moses, Vanderbilt University School of Medicine, Nashville, TN 37232

President-Elect: Lee W. Wattenberg, University of Minnesota, Minneapolis, MN 55455

Treasurer: Thomas J. King, Georgetown University Medical Center, Washington, DC 20007

Executive Director: Margaret Foti, AACR, Public Ledger Building, 620 Chestnut St., Suite 816, Philadelphia, PA 19106-3483

Annual Dues

The annual dues of active members of the American Association for Cancer Research are \$125.00 and they include a subscription to one AACR journal. The member rate for each additional journal is \$45.00. Corresponding members of the Association will be charged an appropriate fee to offset postage costs. Payment of dues and changes of address of members of the Association should be sent promptly to: Member Services, American Association for Cancer Research, Inc., Public Ledger Building, 620 Chestnut St., Suite 816, Philadelphia, PA 19106-3483. Telephone: (215) 440-9300.

Back Issues and Single Copy Sales of the Journal

Copies of back stock of the journal *Cancer Research* may be ordered from Waverly Press, Inc. As long as supplies permit, single copies of *Cancer Research* will be sold by this company at \$30.00 per copy for regular issues and \$30.00 per copy for Supplement issues which contain material from conferences on topics related to cancer. The annual *Proceedings of the American Association for Cancer Research* is available at \$25.00 per copy. Orders outside the U. S. and Canada add \$2.00/copy to offset postage costs.

Advertisements in *Cancer Research*

Advertisement insertion orders and copy must be received 60 days prior to the date of issue in which the advertisement is to be published. The journal is mailed approximately 7 days preceding the date of issue. Inquiries about advertising should be directed to: Elizabeth Mosko, Williams and Wilkins, 428 E. Preston Street, Baltimore, MD 21202-3993. Telephone: (410) 528-4005.

Historical Cover Themes

Readers are invited to submit themes (events, persons, institutions) for consideration for the illustrated covers of *Cancer Research*. Correspondence regarding suggested cover themes, or other matters regarding covers, should be addressed to the Cover Editor, AACR Publications Department.

Submission of Manuscripts

Manuscripts should be sent to the attention of: Dr. Carlo M. Croce, Editor-in-Chief, AACR Publications Department, American Association for Cancer Research, Public Ledger Building, 620 Chestnut St., Suite 816, Philadelphia, PA 19106-3483. Telephone: (215) 440-9300. If accepted, they will be listed under one of ten categories in the Table of Contents. Please specify in a covering letter which category applies to your submission: Basic Sciences section—biochemistry and biophysics, carcinogenesis, endocrinology, immunology, molecular biology and genetics, preclinical pharmacology and experimental therapeutics, tumor biology, virology; Clinical Investigations section—clinical studies, epidemiological studies. (Final categorization in the Table of Contents is at the discretion of the Editor-in-Chief.) Consult the "Instructions for Authors" printed in the January 1 issue of the journal for other submission requirements. Reprints of the "Instructions" are available upon request.

Manuscript Processing Fee

Journal policy requires that a manuscript processing fee of \$75.00 be assessed for each paper to defray the expenses incurred in the editorial review process. Each manuscript submitted for publication should be accompanied by a check (drawn on a U.S. bank) or purchase order for \$75.00 in U. S. currency, payable to *Cancer Research* at the above address. Note: If an author resubmits a manuscript that our Editors previously found unacceptable for publication, it is journal policy to consider it a new submission, assign it a new manuscript number, and charge the author another \$75.00 handling fee to cover the cost of review.

Page Charges

Accepted manuscripts will be published with the understanding that the author(s) will pay a charge of \$65.00 per printed page. Under exceptional circumstances, when no source of grant or other support exists, the author(s) may apply to Dr. Carlo M. Croce, Editor-in-Chief, at the time of submission, for a waiver of the page charges. All such applications must be countersigned by an appropriate institutional official stating that no funds are available for the payment of page charges.

Copyright and Copyright Clearance Center

The Copyright Revision Act (PL 94-553), which became effective January 1, 1978, states that the copyright of a work is vested in the author from the moment of creation. Therefore, all authors who wish to publish in *Cancer Research* must formally transfer copyright to the proprietor of the journal, the American Association for Cancer Research, Inc. It is understood by this transfer that the authors relinquish all exclusive rights of copyright ownership, including the rights of reproduction, derivation, distribution, sale, and display.

Authors who prepared their articles as part of their official duties as employees of the U. S. Federal Government are not required to transfer copyright to the American Association for Cancer Research, Inc., since these articles are considered to be in the public domain. However, it is necessary for these authors to sign the appropriate section of the transfer form. In the case of articles supported by federal grants or contracts, copyright transfer to the American Association for Cancer Research, Inc., is required. The federal government may retain a nonexclusive license to publish or republish such material.

Copies of articles for which the American Association for Cancer Research, Inc. owns the copyright may be made for personal or internal use, provided that the copier pay the per-copy fee of \$2.00 through the Copyright Clearance Center, Inc. This Center is a nonprofit organization through which individuals and institutions may reimburse a copyright owner for photocopying journal articles beyond what is defined as "fair use" in Sections 107 and 108 of the Copyright Revision Act of 1978. Those who wish to photocopy *Cancer Research* articles may report the number of copies they have made, together with the fee code 0008-5472/92 \$02.00, to: Copyright Clearance Center, Inc., 27 Congress St., Salem, MA 01970. Remittances may be sent to the Center at the time of reporting or the Center will bill the user on a monthly basis. Deposit accounts and prepayment plans may also be arranged.

Between June 1978 and August 1983, a fee code appeared on the first page of all articles for which *Cancer Research* owned the copyright. For those issues, it is understood that any articles which did not carry this code are in the public domain.

Cancer Research is abstracted or indexed in *Biological Abstracts*, *Chemical Abstracts*, *Index Medicus*, *Science Citation Index*, and by the *International Cancer Research Data Bank*. This journal is printed on acid-free paper.

No responsibility is accepted by the Editors, by the American Association for Cancer Research, Inc., or by Waverly, Inc. for the opinions expressed by contributors or for the content of the advertisements.

Cancer Research (ISSN 0008-5472) is published twice a month for \$80 per year (for members of the American Association for Cancer Research) or \$350 per year (for nonmembers) by the American Association for Cancer Research, Inc. Second-class postage paid at Baltimore, MD and additional mailing offices. POSTMASTER: Send address changes to *Cancer Research*, c/o Waverly Press, Inc., 428 E. Preston Street, Baltimore, MD 21202-3993.

Copyright 1992 by the American Association for Cancer Research, Inc.